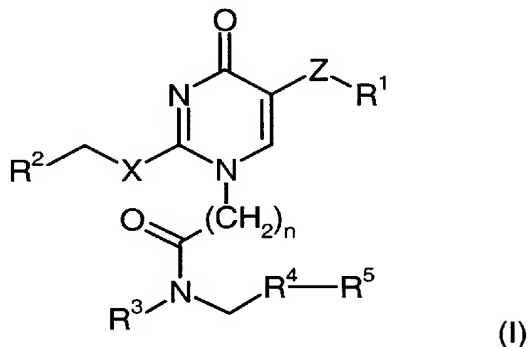


Serial No.: 10/776,876  
Group Art Unit No.: 1624

**Amendments to the Claims:**

1. (Previously Presented) A compound of formula (I):



in which:

$R^1$  is an aryl or heteroaryl group, optionally substituted by 1, 2, 3 or 4 substituents which may be the same or different selected from  $C_{(1-18)}$ alkyl,  $C_{(1-18)}$ alkoxy,  $C_{(1-18)}$ alkylthio, aryl $C_{(1-18)}$ alkoxy, hydroxy, halogen, CN,  $COR^6$ , carboxy,  $COOR^6$ ,  $CONR^9R^{10}$ ,  $NR^6COR^7$ ,  $SO_2NR^9R^{10}$ ,  $NR^6SO_2R^7$ ,  $NR^9R^{10}$ , mono to perfluoro- $C_{(1-4)}$ alkyl and mono to perfluoro- $C_{(1-4)}$ alkoxy, oxo,  $CH_2COOH$  or a salt thereof,  $CH_2COOR^8$ ,  $CH_2CONR^9R^{10}$ ,  $CH_2CN$ ,  $(CH_2)_mNR^9R^{10}$ ,  $(CH_2)_mOH$  or  $(CH_2)_mOR^6$  where m is an integer from 1 to 3;

$R^2$  is an aryl or heteroaryl group, optionally substituted by 1, 2, 3 or 4 substituents which may be the same or different selected from  $C_{(1-18)}$ alkyl,  $C_{(1-18)}$ alkoxy,  $C_{(1-18)}$ alkylthio, aryl $C_{(1-18)}$ alkoxy, hydroxy, halogen, CN,  $COR^6$ , carboxy,  $COOR^6$ ,  $CONR^9R^{10}$ ,  $NR^6COR^7$ ,  $SO_2NR^9R^{10}$ ,  $NR^6SO_2R^7$ ,  $NR^9R^{10}$ , mono to perfluoro- $C_{(1-4)}$ alkyl, mono to perfluoro- $C_{(1-4)}$ alkoxy, and aryl $C_{(1-4)}$ alkyl;

$R^3$  is hydrogen or  $C_{(1-4)}$ alkyl which may be unsubstituted or substituted by hydroxy,  $OR^6$ ,  $COR^6$ , carboxy,  $COOR^6$ ,  $CONR^9R^{10}$ ,  $NR^9R^{10}$ , mono- or di-(hydroxy $C_{(1-6)}$ alkyl)amino or N-hydroxy $C_{(1-6)}$ alkyl-N- $C_{(1-6)}$ alkyl amino;

$R^4$  is an aryl or a heteroaryl ring optionally substituted by 1, 2, 3 or 4 substituents which may be the same or different selected from  $C_{(1-18)}$ alkyl,  $C_{(1-18)}$ alkoxy,  $C_{(1-18)}$ alkylthio, aryl $C_{(1-18)}$ alkoxy, hydroxy, halogen, CN,  $COR^6$ , carboxy,  $COOR^6$ ,  $CONR^9R^{10}$ ,  $NR^6COR^7$ ,  $SO_2NR^9R^{10}$ ,  $NR^6SO_2R^7$ ,  $NR^9R^{10}$ , mono to perfluoro- $C_{(1-4)}$ alkyl and mono to perfluoro- $C_{(1-4)}$ alkoxy;

Serial No.: 10/776,876  
Group Art Unit No.: 1624

R<sup>5</sup> is an aryl ring which is further optionally substituted by 1, 2, 3 or 4 substituents which may be the same or different selected from C<sub>(1-18)</sub>alkyl, C<sub>(1-18)</sub>alkoxy, C<sub>(1-18)</sub>alkylthio, arylC<sub>(1-18)</sub>alkoxy, hydroxy, halogen, CN, COR<sup>6</sup>, carboxy, COOR<sup>6</sup>, CONR<sup>9</sup>R<sup>10</sup>, NR<sup>6</sup>COR<sup>7</sup>, SO<sub>2</sub>NR<sup>9</sup>R<sup>10</sup>, NR<sup>6</sup>SO<sub>2</sub>R<sup>7</sup>, NR<sup>9</sup>R<sup>10</sup>, mono to perfluoro-C<sub>(1-4)</sub>alkyl and mono to perfluoro-C<sub>(1-4)</sub>alkoxy;

R<sup>6</sup> and R<sup>7</sup> are independently hydrogen or C<sub>(1-20)</sub>alkyl, for instance C<sub>(1-4)</sub>alkyl (e.g. methyl or ethyl);

R<sup>8</sup> is C<sub>(1-4)</sub>alkyl or a pharmaceutically acceptable *in vivo* hydrolysable ester group;

R<sup>9</sup> and R<sup>10</sup> which may be the same or different is each selected from hydrogen, C<sub>(1-12)</sub>alkyl, CH<sub>2</sub>R<sup>11</sup>, CHR<sup>12</sup>CO<sub>2</sub>H or a salt thereof, or R<sup>9</sup> and R<sup>10</sup> together with the nitrogen to which they are attached form a 4- to 7-, preferably 5- to 7-, membered ring optionally containing one or more further heteroatoms selected from oxygen, nitrogen and sulphur, and optionally substituted by one or two substituents selected from hydroxy, oxo, C<sub>(1-4)</sub>alkyl, C<sub>(1-4)</sub>alkylCO, or aryl;

R<sup>11</sup> is COOH or a salt thereof, COOR<sup>8</sup>, CONR<sup>6</sup>R<sup>7</sup>, CN, CH<sub>2</sub>OH or CH<sub>2</sub>OR<sup>6</sup>;

R<sup>12</sup> is an amino acid side chain;

n is an integer from 1 to 4, preferably 1 or 3;

X is O or S; and

Z is CR<sup>13</sup>R<sup>14</sup> where R<sup>13</sup> and R<sup>14</sup> are each hydrogen or C<sub>(1-4)</sub>alkyl, or R<sup>13</sup> and R<sup>14</sup> together with the intervening carbon atom form a C<sub>(3-6)</sub>cycloalkyl ring.

2. (Original) A compound of formula (I) as claimed in claim 1 in which Z is CH<sub>2</sub>.

3. (Previously Presented) A compound of formula (I) as claimed in claim 1 in which R<sup>1</sup> is an aryl group selected from phenyl and naphthyl or a heteroaryl group which comprises a 5- or 6- membered, monocyclic heteroaryl group comprising 1 or 2 nitrogen heteroatoms.

4. (Previously Presented) A compound of formula (I) as claimed in claim 1 in which R<sup>1</sup> is pyrimidyl optionally substituted by 1 or 2 substituents selected from oxo, arylC<sub>(1-4)</sub>alkyl, C<sub>(1-6)</sub>alkyl, C<sub>(3-6)</sub>cycloalkyl, hydroxy, C<sub>(1-4)</sub>alkoxy,

Serial No.: 10/776,876  
Group Art Unit No.: 1624

carboxyC<sub>(1-6)</sub>alkyl, C<sub>(1-6)</sub>alkylcarboxyC<sub>(1-6)</sub>alkyl, di-C<sub>(1-6)</sub>alkylamino, and morpholino; or pyrazolyl optionally substituted by C<sub>(1-6)</sub>alkyl.

5. (Original) A compound as claimed in claim 4 in which ZR<sup>1</sup> is pyrimid-5-ylmethyl optionally substituted by 2-methoxy, 2-trifluoromethyl, 2-(4-morpholino) or 2-dimethylamino; 2-oxo-pyrimid-5-ylmethyl or 1-methyl-4-pyrazolylmethyl.

6. (Previously Presented) A compound of formula (I) as claimed in claim 1 in which X is S.

7. (Previously Presented) A compound of formula (I) as claimed in claim 1 in which R<sup>2</sup> is an aryl group selected from phenyl and naphthyl or a heteroaryl group selected from pyridyl, pyrimidinyl, pyrazolyl, furanyl, thienyl, thiazolyl, quinolyl, benzothiazolyl, pyridazolyl and pyrazinyl.

8. (Original) A compound of formula (I) as claimed in claim 7 in which R<sup>2</sup> is phenyl optionally substituted by halogen

9. (Previously Presented) A compound of formula (I) as claimed in claim 1 in which R<sup>3</sup> is selected from hydrogen; and methyl, ethyl and propyl, optionally substituted by amino, C<sub>(1-3)</sub>alkylamino, di C<sub>(1-3)</sub>alkylamino, hydroxyC<sub>(1-3)</sub>alkylamino, hydroxy, C<sub>(1-3)</sub>alkoxy, carboxy, C<sub>(1-3)</sub>alkylcarboxy, and heterocycyl.

10. (Previously Presented) A compound of formula (I) as claimed in claim 1 in which R<sup>4</sup> is selected from phenyl optionally substituted by halogen; thiophene; pyridine; and pyrimidine.

11. (Previously Presented) A compound of formula (I) as claimed in claim 1 in which R<sup>5</sup> is phenyl optionally substituted by halogen, trifluoromethyl, or trifluoromethoxy.

Serial No.: 10/776,876  
Group Art Unit No.: 1624

12. (Previously Presented) A compound of formula (I) as claimed in claim 10 in which R<sup>4</sup> and R<sup>5</sup> together form a 4-(phenyl)phenyl substituent in which the remote phenyl ring may be optionally substituted by halogen or trifluoromethyl.

13. (Cancelled).

14. (Previously presented) A compound of formula (I) as claimed in claim 1 selected from the group consisting of:

1-(N-methyl-N-(4-(4-chlorophenyl)benzyl)aminocarbonylmethyl)-2-(4-fluorobenzyl)thio-5-(1-methylpyrazol-4-ylmethyl)pyrimidin-4-one;

1-(N-methyl-N-(4-(4-trifluoromethylphenyl)benzyl)aminocarbonylmethyl)-2-(4-fluorobenzyl)thio-5-(1-methylpyrazol-4-ylmethyl)pyrimidin-4-one;

1-(N-(2-dimethylaminoethyl)-N-(4-(4-chlorophenyl)benzyl)aminocarbonylmethyl)-2-(4-fluorobenzyl)thio-5-(1-methylpyrazol-4-ylmethyl)pyrimidin-4-one;

1-(N-methyl-N-(4-(4-chlorophenyl)benzyl)aminocarbonylmethyl)-2-(4-fluorobenzyl)thio-5-(2-(4-morpholino)pyrimid-5-ylmethyl)pyrimidin-4-one;

1-(N-(2-(dimethylamino)ethyl)-N-(4-(4-trifluoromethylphenyl)benzyl)aminocarbonylmethyl)-2-(4-fluorobenzyl)thio-5-(1-methyl-4-pyrazolylmethyl)pyrimidin-4-one;

1-(N-(2-(diethylamino)ethyl)-N-(4-(4-chlorophenyl)benzyl)aminocarbonylmethyl)-2-(4-fluorobenzyl)thio-5-(1-methyl-4-pyrazolylmethyl)pyrimidin-4-one;

1-(N-(2-(diethylamino)ethyl)-N-(2-(4-trifluoromethylphenyl)pyrid-5-ylmethyl)aminocarbonylmethyl)-2-(4-fluorobenzyl)thio-5-(1-methyl-4-pyrazolylmethyl)pyrimidin-4-one;

1-(N-(2-(1-piperidino)ethyl)-N-(4-(4-trifluoromethylphenyl)benzyl)aminocarbonylmethyl)-2-(4-fluorobenzyl)thio-5-(1-methyl-4-pyrazolylmethyl)pyrimidin-4-one bitartrate;

1-(N-(carboxymethyl)-N-(4-(4-trifluoromethylphenyl)benzyl)aminocarbonylmethyl)-2-(4-fluorobenzyl)thio-5-(1-methyl-4-pyrazolylmethyl)pyrimidin-4-one sodium salt; or

a pharmaceutically acceptable salt thereof.

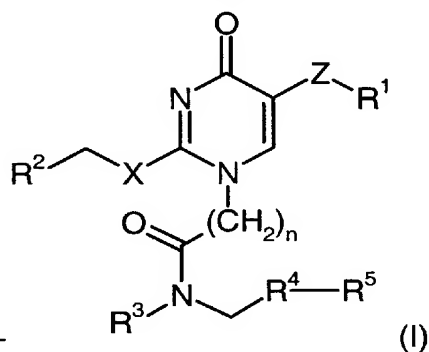
Serial No.: 10/776,876  
Group Art Unit No.: 1624

15. (Previously Presented) A pharmaceutical composition comprising a compound of formula (I) as claimed in claim 14 and a pharmaceutically acceptable carrier.

16. – 18. (Cancelled)

19. (Original) A method of treating atherosclerosis which method comprises administering to a patient in need thereof an effective amount of a compound of formula (I) as claimed in claim 1 and a statin.

20. (Currently Amended) A process for preparing a compound of formula (I) as



~~defined in claim 1~~

in which:

R<sup>1</sup> is an aryl or heteroaryl group, optionally substituted by 1, 2, 3 or 4 substituents which may be the same or different selected from C<sub>(1-18)</sub>alkyl, C<sub>(1-18)</sub>alkoxy, C<sub>(1-18)</sub>alkylthio, arylC<sub>(1-18)</sub>alkoxy, hydroxy, halogen, CN, COR<sup>6</sup>, carboxy, COOR<sup>6</sup>, CONR<sup>9</sup>R<sup>10</sup>, NR<sup>6</sup>COR<sup>7</sup>, SO<sub>2</sub>NR<sup>9</sup>R<sup>10</sup>, NR<sup>6</sup>SO<sub>2</sub>R<sup>7</sup>, NR<sup>9</sup>R<sup>10</sup>, mono to perfluoro-C<sub>(1-4)</sub>alkyl and mono to perfluoro-C<sub>(1-4)</sub>alkoxy, oxo, CH<sub>2</sub>COOH or a salt thereof, CH<sub>2</sub>COOR<sup>8</sup>, CH<sub>2</sub>CONR<sup>9</sup>R<sup>10</sup>, CH<sub>2</sub>CN, (CH<sub>2</sub>)<sub>m</sub>NR<sup>9</sup>R<sup>10</sup>, (CH<sub>2</sub>)<sub>m</sub>OH or (CH<sub>2</sub>)<sub>m</sub>OR<sup>6</sup> where m is an integer from 1 to 3;

R<sup>2</sup> is an aryl or heteroaryl group, optionally substituted by 1, 2, 3 or 4 substituents which may be the same or different selected from C<sub>(1-18)</sub>alkyl, C<sub>(1-18)</sub>alkoxy, C<sub>(1-18)</sub>alkylthio, arylC<sub>(1-18)</sub>alkoxy, hydroxy, halogen, CN, COR<sup>6</sup>, carboxy, COOR<sup>6</sup>, CONR<sup>9</sup>R<sup>10</sup>, NR<sup>6</sup>COR<sup>7</sup>, SO<sub>2</sub>NR<sup>9</sup>R<sup>10</sup>, NR<sup>6</sup>SO<sub>2</sub>R<sup>7</sup>, NR<sup>9</sup>R<sup>10</sup>, mono to perfluoro-C<sub>(1-4)</sub>alkyl, mono to perfluoro-C<sub>(1-4)</sub>alkoxy, and arylC<sub>(1-4)</sub>alkyl;

R<sup>3</sup> is hydrogen or C<sub>(1-4)</sub>alkyl which may be unsubstituted or substituted by hydroxy, OR<sup>6</sup>, COR<sup>6</sup>, carboxy, COOR<sup>6</sup>, CONR<sup>9</sup>R<sup>10</sup>, NR<sup>9</sup>R<sup>10</sup>, mono- or di-(hydroxyC<sub>(1-6)</sub>alkyl)amino or N-hydroxyC<sub>(1-6)</sub>alkyl-N-C<sub>(1-6)</sub>alkyl amino;

Serial No.: 10/776,876  
Group Art Unit No.: 1624

R<sup>4</sup> is an aryl or a heteroaryl ring optionally substituted by 1, 2, 3 or 4 substituents which may be the same or different selected from C<sub>(1-18)</sub>alkyl, C<sub>(1-18)</sub>alkoxy, C<sub>(1-18)</sub>alkylthio, arylC<sub>(1-18)</sub>alkoxy, hydroxy, halogen, CN, COR<sup>6</sup>, carboxy, COOR<sup>6</sup>, CONR<sup>9</sup>R<sup>10</sup>, NR<sup>6</sup>COR<sup>7</sup>, SO<sub>2</sub>NR<sup>9</sup>R<sup>10</sup>, NR<sup>6</sup>SO<sub>2</sub>R<sup>7</sup>, NR<sup>9</sup>R<sup>10</sup>, mono to perfluoro-C<sub>(1-4)</sub>alkyl and mono to perfluoro-C<sub>(1-4)</sub>alkoxy;

R<sup>5</sup> is an aryl ring which is further optionally substituted by 1, 2, 3 or 4 substituents which may be the same or different selected from C<sub>(1-18)</sub>alkyl, C<sub>(1-18)</sub>alkoxy, C<sub>(1-18)</sub>alkylthio, arylC<sub>(1-18)</sub>alkoxy, hydroxy, halogen, CN, COR<sup>6</sup>, carboxy, COOR<sup>6</sup>, CONR<sup>9</sup>R<sup>10</sup>, NR<sup>6</sup>COR<sup>7</sup>, SO<sub>2</sub>NR<sup>9</sup>R<sup>10</sup>, NR<sup>6</sup>SO<sub>2</sub>R<sup>7</sup>, NR<sup>9</sup>R<sup>10</sup>, mono to perfluoro-C<sub>(1-4)</sub>alkyl and mono to perfluoro-C<sub>(1-4)</sub>alkoxy;

R<sup>6</sup> and R<sup>7</sup> are independently hydrogen or C<sub>(1-20)</sub>alkyl, for instance C<sub>(1-4)</sub>alkyl (e.g. methyl or ethyl);

R<sup>8</sup> is C<sub>(1-4)</sub>alkyl or a pharmaceutically acceptable *in vivo* hydrolysable ester group;

R<sup>9</sup> and R<sup>10</sup> which may be the same or different is each selected from hydrogen, C<sub>(1-12)</sub>alkyl, CH<sub>2</sub>R<sup>11</sup>, CHR<sup>12</sup>CO<sub>2</sub>H or a salt thereof, or R<sup>9</sup> and R<sup>10</sup> together with the nitrogen to which they are attached form a 4- to 7-, preferably 5- to 7-, membered ring optionally containing one or more further heteroatoms selected from oxygen, nitrogen and sulphur, and optionally substituted by one or two substituents selected from hydroxy, oxo, C<sub>(1-4)</sub>alkyl, C<sub>(1-4)</sub>alkylCO, or aryl;

R<sup>11</sup> is COOH or a salt thereof, COOR<sup>8</sup>, CONR<sup>6</sup>R<sup>7</sup>, CN, CH<sub>2</sub>OH or CH<sub>2</sub>OR<sup>6</sup>;

R<sup>12</sup> is an amino acid side chain;

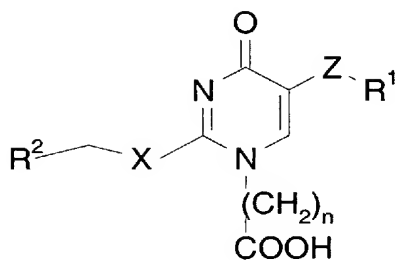
n is an integer from 1 to 4, preferably 1 or 3;

X is O or S; and

Z is CR<sup>13</sup>R<sup>14</sup> where R<sup>13</sup> and R<sup>14</sup> are each hydrogen or C<sub>(1-4)</sub>alkyl, or R<sup>13</sup> and R<sup>14</sup> together with the intervening carbon atom form a C<sub>(3-6)</sub>cycloalkyl ring;  
which process comprises:

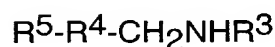
(a) reacting a compound of formula (II):

Serial No.: 10/776,876  
Group Art Unit No.: 1624



(II)

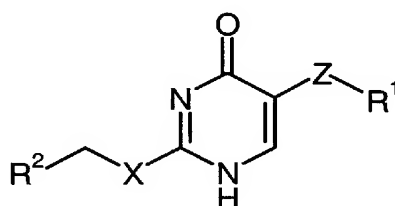
in which X, Y, Z, R<sup>1</sup> and R<sup>2</sup> are the same as defined herein above for formula (I) as defined in claim 1,  
with a compound of formula (III):



(III)

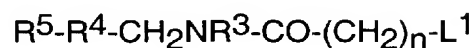
in which R<sup>3</sup>, R<sup>4</sup> and R<sup>5</sup> are the same as defined herein above for formula (I) as defined in claim 1; under amide forming conditions;

(b) reacting a compound of formula (IV):



(IV)

in which X, Z, R<sup>1</sup> and R<sup>2</sup> are the same as defined herein above for formula (I) as defined in claim 1, with a compound of formula (V):

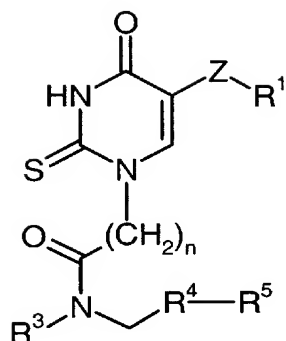


(V)

in which n, R<sup>3</sup>, R<sup>4</sup> and R<sup>5</sup> are the same as defined herein above for formula (I) as defined in claim 1, and L<sup>1</sup> is a leaving group such as halogen; in the presence of a base such as a secondary or tertiary amine, in an inert solvent;

(c) when X is S, reacting a compound of formula (VI):

Serial No.: 10/776,876  
Group Art Unit No.: 1624



(VI)

in which n, Z, R<sup>1</sup>, R<sup>3</sup>, R<sup>4</sup> and R<sup>5</sup> are the same as defined herein above for formula (I) ~~as defined in claim 1~~, with a compound of formula (VII):

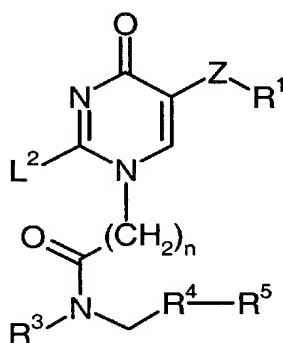


(VII)

in which R<sup>2</sup> and L<sup>1</sup> are the same as defined herein above for formula (I) ~~as defined in claim 1~~,

in the presence of a base such as a secondary or tertiary amine, in an inert solvent;  
or

(d) when X is O, reacting a compound of formula (VIII):

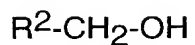


(VIII)

in which n, Z, R<sup>1</sup>, R<sup>3</sup>, R<sup>4</sup> and R<sup>5</sup> are the same as defined herein above for formula (I) ~~as defined in claim 1~~, and L<sup>2</sup> is a leaving group,  
with a compound of formula (IX):



Serial No.: 10/776,876  
Group Art Unit No.: 1624



(IX)

in which  $R^2$  is the same as defined herein above for formula (I) as defined in claim 1,  
in the presence of a base, in an inert solvent.

21. (Previously Presented) A pharmaceutical composition comprising a compound of formula (I) as claimed in claim 1 and a pharmaceutically acceptable carrier.

22. (Previously Presented) A method of treating atherosclerosis which method comprises administering to a patient in need thereof an effective amount of a compound of formula (I) as claimed in claim 1 to a patient in need thereof.